



PATENTS

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:
Balaji Venkataraman

Application No.: **10/635,428**

Filed: **August 6, 2003**

For: **VITAMIN COMPOSITIONS FOR THE
TREATMENT AND PREVENTION OF
VASCULAR DISEASE AND DEMENTIA**

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)
) Examiner: **Not assigned**

)
) Art Unit: **Not assigned**
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INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Sir:

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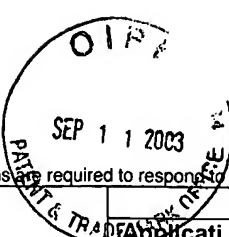
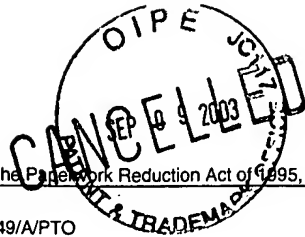
Respectfully submitted,

Lisa C. Pavento
Reg. No. 44,669

KILPATRICK STOCKTON LLP
1100 Peachtree Street
Suite 2800
Atlanta, Georgia 30309-4530
(404) 815-6500
Our Docket: 52761-0100 (285976)

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on September 9, 2003.

Lisa C. Pavento - Reg. No. 44,669



INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)		Complete if Known	
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		Group Art Unit	
Examiner Name			
Attorney Docket Number	52761/285976		

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U.S. PATENT DOCUMENTS

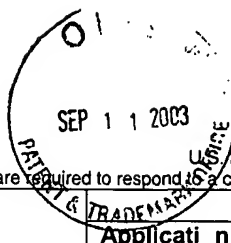
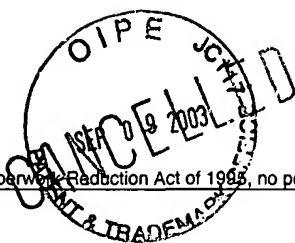
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		Number	Kind Code ² (if known)			
	1.	5,494,678		Paradissis, <i>et al.</i>	02-27-1996	
	2.	5,563,126		Allen, <i>et al.</i>	10-08-1996	
	3.	5,795,873		Allen	08-18-1998	
	4.	5,801,161		Merkus	09-01-1998	
	5.	5,932,624		Herbert	08-03-1999	
	6.	5,962,048		Register	10-05-1999	
	7.	6,039,978		Bangs, <i>et al.</i>	03-21-2000	
	8.	6,048,846		Cochran	04-11-2000	
	9.	6,121,249		Weissman, <i>et al.</i>	09-19-2000	
	10.	6,127,370		Smith, <i>et al.</i>	10-03-2000	
	11.	6,207,651	B1	Allen, <i>et al.</i>	03-27-2001	
	12.	6,299,896	B1	Cooper, <i>et al.</i>	10-09-2001	
	13.	6,323,188	B1	Weissman	11-27-2001	
	14.	6,361,800	B1	Cooper, <i>et al.</i>	03-26-2002	
	15.	6,369,041	B2	Horrobin, <i>et al.</i>	04-09-2002	
	16.	6,479,545	B1	Levinson, <i>et al.</i>	11/12-2002	
	17.	6,528,496	B1	Allen, <i>et al.</i>	03-04-2003	
	18.	2002/0010148	A1	Horrobin, <i>et al.</i>	01-24-2002	
	19.	2002/0136711	A1	Cochran	09-26-2002	
	20.	2002/0146463	A1	Clayton	10-10-2002	
	21.	2003/0049352	A1	Mehansho, <i>et al.</i>	03-13-2003	
	22.	2003/017136 (Abstract)	A1	Cruz, <i>et al.</i>	01-23-2003	

FOREIGN PATENT DOCUMENTS

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		Office ³	Number ⁴	Kind Code ⁵ (if known)				
	23.	EP	0366480 (Abstract)	B1	Bar Ilan University	08-24-1994		
	24.	WO	96/11009 (Abstract)	A1	UK	04-18-1996		
	25.	WO	98/01157 (Abstract)	A1	WWK Trust, Worsley, Andrew Peter	01-15-1998		
	26.	WO	2000/056295 (Abstract)	A2	Kilgowan Limited	09-28-2000		
	27.	WO	2001/056609 (Abstract)	A1	Germany	08-09-2001		
	28.	WO	2002/087593 (Abstract)	A1	Cobalz Limited	11-07-2002		

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**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**

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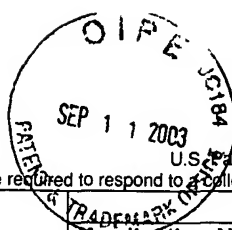
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	29.	ALTURA, BURTON M. and ALTURA, BELLA T., "Magnesium and Cardiovascular Biology: An Important Link Between Cardiovascular Risk Factors and Atherogenesis," Cellular and Molecular Biology Research, 1995, pp. 347-359, Vol. 41, No. 5.	
	30.	ANDERSSON, H.C., <i>et al.</i> , "Long-Term Outcome in Treated Combined Methylmalonic Acidemia and Homocystinemia," Genet Med, May-June 1999, pp. 146-150. (Abstract)	
	31.	APPEL, LAWRENCE, J., <i>et al.</i> , "A Clinical Trial of the Effects of Dietary Patterns on Blood Pressure," N. Engl. J. Med, April 17, 1997, pp. 1117-1124, Vol. 336, No. 16.	
	32.	BASU, TAPAN K. and MANN, SARABJIT, "Vitamin B-6 Normalizes the Altered Sulfur Amino Acid Status of Rats Fed Diets Containing Pharmacological Levels of Niacin Without Reducing Niacin's Hypolipidemic Effects," J. Nutr., 1997, pp.117-121, Vol. 127.	
	33.	BLOOM, O. JOSH, <i>et al.</i> , "Do Antioxidants (Vitamins C, E) Improve Outcomes in Patients With Coronary Artery Disease?" J. Fam. Prac., November 2002, p. 978, Vol. 51, No. 11.	
	34.	BLUMBERG, A., <i>et al.</i> , "Vitamin Nutrition in Patients on Continuous Ambulatory Peritoneal Dialysis (CAPD)," Clin Nephrol, November 1983, pp. 244-250, Vol. 20, No. 5. (Abstract)	
	35.	BOUSHEY, CAROL J., <i>et al.</i> , "A Quantitative Assessment of Plasma Homocysteine as a Risk Factor for Vascular Disease," JAMA, October 4, 1995, pp. 1049-1057, Vol. 274, No. 13.	
	36.	BRATTSTROM, L. E., <i>et al.</i> , "Folic Acid - An Innocuous Means to Reduce Plasma Homocysteine," Scand J. Clin. Lab Invest., 1998, pp. 215-221, Vol. 48.	
	37.	BRATTSTROM, LARS E., <i>et al.</i> , "Folic Acid Responsive Postmenopausal Homocysteinemia, Metabolism," November 1985, pp. 1073-1077, Vol. 34, No. 11.	
	38.	BRATTSTROM, LARS, <i>et al.</i> , "Impaired Homocysteine Metabolism in Early-Onset Cerebral and Peripheral Occlusive Arterial Disease," Atherosclerosis, 1990, pp. 51-60, Vol. 81.	
	39.	CAMPBELL, BRUCE G., <i>et al.</i> , "Do Calcium Supplements Prevent Postmenopausal Osteoporotic Fractures?" J of Fam Prac., March 2003, pp. 236-237, Vol. 52, No. 3.	
	40.	CHAMBERS, JOHN C., <i>et al.</i> , "Improved Vascular Endothelial Function After Oral B Vitamins," Circulation, Nov. 14, 2000, pp. 2479-2483, Vol. 102.	

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¹Unique citation designation number. ²Applicant is to place a check mark here if English language translation is attached.



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OTHER INFORMATION - NON PATENT LITERATURE DOCUMENTS

Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T ²
	41.	CHAN, P., <i>et al.</i> , "Randomized, Double-Blind, Placebo-Controlled Study of the Safety and Efficacy of Vitamin B Complex in the Treatment of Nocturnal Leg Cramps in Elderly Patients with Hypertension," J Clin Pharmacol., 1998, pp. 1151-1154, Vol. 38, No. 12. (Abstract)	
	42.	DOSHI, SAGAR N., <i>et al.</i> , "Folic Acid Improves Endothelial Function in Coronary Artery Disease via Mechanisms Largely Independent of Homocysteine Lowering," Circulation, January 1/8, 2002, pp. 23-26, Vol. 102.	
	43.	DUDMAN, NICHOLAS P. B., <i>et al.</i> , "Homocysteinemia: Depressed Plasma Serine Levels," Metabolism, February 1987, pp. 198-201, Vol. 36, No. 2.	
	44.	ELIAN, <i>et al.</i> , "Hydroxocobalamin Reduces Hyperhomocysteinemia in End-Stage Renal Disease," Metabolism, July 2002, pp. 881-886, Vol. 51. (Abstract)	
	45.	FANG, JAMES C., <i>et al.</i> , "Effect of Vitamins C and E on Progression of Transplant-Associated Arteriosclerosis: A Randomised Trial," The Lancet, March 30, 2002, pp. 1108-1113, Vol. 359.	
	46.	FRANKEN, D.G., <i>et al.</i> , "Effect of Various Regimens of Vitamin B ₆ and Folic Acid on Mild Hyperhomocysteinaemia in Vascular Patients," J. Inher. Metab. Dis., 1994, pp. 159-162, Vol. 17.	
	47.	GABY, A. R., "Intravenous Nutrient Therapy: The Myers' Cocktail," Altern Med Rev, Oct. 2002, pp. 389-403.	
	48.	GARG, REKKA, <i>et al.</i> , "Niacin Treatment Increases Plasma Homocyst(e)ine Levels," Am Heart J., December 1999, pp 1082-1087, Vol. 138, No.6, Part 1.	
	49.	GOTTFRIES, C. G., "Pharmacological Treatment Strategies in Alzheimer Type Dementia," Eur Neuropsychopharmacol, Nov. 1990, pp. 1-5. (Abstract)	
	50.	HAIDARI, MEHRAN, <i>et al.</i> , "Enhanced Susceptibility to Oxidation and Diminished Vitamin E Content of LDL from Patients with Stable Coronary Artery Disease," Clinical Chemistry, 2001, pp. 1234-1240, Vol. 47, No.7.	
	51.	HARPEY, JEAN-PAUL, <i>et al.</i> , "Homocystinuria Caused by 5,10-Methylenetetrahydrofolate Reductase Deficiency: A Case in an Infant Responding to Methionine, Folinic Acid, Pyridoxine, and Vitamin B ₁₂ Therapy," J Pediatrics, February 1981, pp. 275-278, Vol. 98, No. 2.	

Examiner Signature		Date Considered	
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	52.	HEINECKE, JAY W., "Is the Emperor Wearing Clothes?" Arterioscler Thromb Vasc Biol., August 2001, pp. 1261-1264, Vol. 21.	
	53.	HEINONEN, O.P., <i>et al.</i> , "The Effect of Vitamin E and Beta Carotene on the Incidence of Lung Cancer and Other Cancers in Male Smokers," TN Engl J Med, April 14, 1994, pp. 1029-1035, Vol. 330, No. 15.	
	54.	HENKIN, YAAKOV, <i>et al.</i> , "Niacin Revisited: Clinical Observations on an Important But Underutilized Drug," Am J Med, September 1991, pp. 239-246, Vol. 91.	
	55.	HERBERT, V., "The Inhibition and Promotion of Cancers by Folic Acid, Vitamin B ₁₂ , and their Antagonists," American Chemical Society (ACS Symposium series), 1985, pp. 32-36, Vol. 277.	
	56.	KAUL, NALINI, <i>et al.</i> , "α-Tocopherol and Atherosclerosis," E.B.M., 2001, pp. 5-12, Vol. 226.	
	57.	KLERK, M., "Effect of Homocysteine Reduction by B-Vitamin Supplementation on Markers of Clotting Activation," Thromb Haemost, Aug. 2002, pp. 230-235, Vol. 88.	
	58.	LIAO, FANGZI, <i>et al.</i> , "Is Low Magnesium Concentration a Risk Factor For Coronary Heart Disease? The Atherosclerosis Risk in Communities (ARIC) Study," Am Heart J, September 1998, pp. 480-490, Vol. 136, No. 3.	
	59.	MARCHESONI, D., <i>et al.</i> , "Menopause Rather than Estrogen Modifies Plasma Homocysteine Levels," International Journal of Gynecology and Obstetrics, 2003, pp. 293-297, Vol. 81.	
	60.	MASON, JB and MILLER, J.W., "The Effects of Vitamins B12, B6, and Folate on Blood Homocysteine Levels," Ann NY Acad Sci., Sept. 30, 1992, pp. 197-203.	
	61.	NILSSON, K., <i>et al.</i> , "Treatment of Cobalamin Deficiency in Dementia, Evaluated Clinically and With Cerebral Blood Flow Measurements," Aging (Milano), June 2000, pp. 199-207, Vol. 12.	
	62.	OLSZEWSKI, ANDRZEJ J., <i>et al.</i> , "Reduction of Plasma Lipid and Homocysteine Levels by Pyridoxine, Folate, Cobalamin, Choline, Riboflavin, and Troxerutin in Atherosclerosis," Atherosclerosis, 1989, pp. 1-6, Vol. 75.	
	63.	ROSSI, A., <i>et al.</i> , "Early-Onset Combined Methymalonic Aciduria and Homocystinuria: Neuroradiologic Findings," Am J. Neuroradiol, March 2001, pp. 554-563, Vol. 22.	

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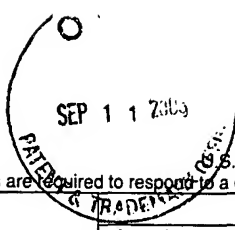
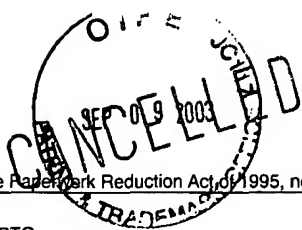
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	64.	ROSSOUW, JACQUES E., <i>et al.</i> , "Risks and Benefits of Estrogen Plus Progestin in Healthy Postmenopausal Women" JAMA, July 17, 2002, pp. 321-333, Vol. 288, No. 3.	
	65.	SCHNYDER, GUIDO, <i>et al.</i> , "Effect of Homocysteine-Lowering Therapy with Folic Acid, Vitamin B ₁₂ , and Vitamin B ₆ on Clinical Outcome After Percutaneous Coronary Intervention," JAMA, August 28, 2002, pp. 973-979, Vol. 288, No. 8.	
	66.	SHECHTER, MICHAEL, <i>et al.</i> , "Oral Magnesium Supplementation Inhibits Platelet-Dependent Thrombosis in Patients With Coronary Artery Disease," Am J Cardio., July 15, 1999, pp. 152-156, Vol. 84.	
	67.	SHECHTER, MICHAEL, <i>et al.</i> , "Oral Magnesium Therapy Improves Endothelial Function in Patients with Coronary Artery Disease," Circulation, November 7, 2000, pp. 2351-2358, Vol. 102.	
	68.	STABLER, <i>et al.</i> , "Elevations of Serum Cystathionine and Homocysteine in Vitamin B-6, Folate and Cobalamin Deficient Rats," Blood, 1994, pp. 118A, Vol 84, No. 10, Suppl. 1.	
	69.	STAMPER, MEIR J. and MALINOW, M. RENE, "Can Lowering Homocysteine Levels Reduce Cardiovascular Risk?" The New England Journal of Medicine, Feb. 2, 1995, pp. 328-329, Vol. 332, No. 5.	
	70.	STEIN, JAMES H. and MCBRIDE, PATRICK E., "Hyperhomocysteinemia and Atherosclerotic Vascular Disease," Arch Intern Med., June 22, 1998; pp. 1301-1306, Vol. 158.	
	71.	STEPHENS, NIGEL G., <i>et al.</i> , "Randomised Controlled Trial of Vitamin E in Patients with Coronary Disease: Cambridge Heart Antioxidant Study (CHAOS)," The Lancet, March 23, 1996, pp. 781-786, Vol. 347.	
	72.	STIPANUK, MARTHA H., "Metabolism of Sulfur-Containing Amino Acids," Ann. Rev. Nutr., 1986, pp. 179-209, Vol. 6.	
	73.	TICE, JEFFREY A., <i>et al.</i> , "Cost-effectiveness of Vitamin Therapy to Lower Plasma Homocysteine Levels for the Prevention of Coronary Heart Disease: Effect of Grain Fortification and Beyond," JAMA, August 22/29, 2001, pp. 936-943, Vol. 286.	

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	74.	UELAND, P.M. and REFSUM, H., "Plasma Homocysteine, A Risk Factor For Premature Vascular Disease. Plasma Levels in Healthy Person; During Pathologic Conditions and Drug Therapy," Nord Med., 1989, pp. 293-308, Vol. 104, No. 11. (Abstract)	
	75.	VALAGUSSA, F., <i>et al.</i> , "Dietary Supplementation With N-3 Polyunsaturated Fatty Acids and Vitamin E After Myocardial Infarction: Results of the GISSI-Prevenzione Trial," The Lancet, August 7, 1999, pp. 447-455, Vol. 354.	
	76.	VAN DER KUY, <i>et al.</i> , "Hydroxocobalamin, A Nitric Oxide Scavenger, In the Prophylaxis of Migraine: An Open, Pilot Study," Cephalalgia Sept. 2002, pp. 513-519, Vol. 22, No. 7,.	
	77.	VERHAAR, MARIANNE C., <i>et al.</i> , "Effects of Oral Folic Acid Supplementation on Endothelial Function in Familial Hypercholesterolemia," Circulation, July 27, 1999, pp. 335-338, Vol. 100.	
	78.	VIVANCO, N., <i>et al.</i> , "Treatment of Anorexia Nervosa," Rev Med Chil, 1992, pp. 69-75, Vol. 120. (Abstract)	
	79.	WATERS, DAVID D., <i>et al.</i> , "Effects of Hormone Replacement Therapy and Antioxidant Vitamin Supplements on Coronary Atherosclerosis in Postmenopausal Women," JAMA, November 20, 2002, pages 2432-3440, Vol 288, No. 19.	
	80.	YANG, CHUN-YUH, "Calcium and Magnesium in Drinking Water and Risk of Death from Cerebrovascular Disease," Stroke, 1998, pp. 411-414, Vol. 29.	

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